

#5

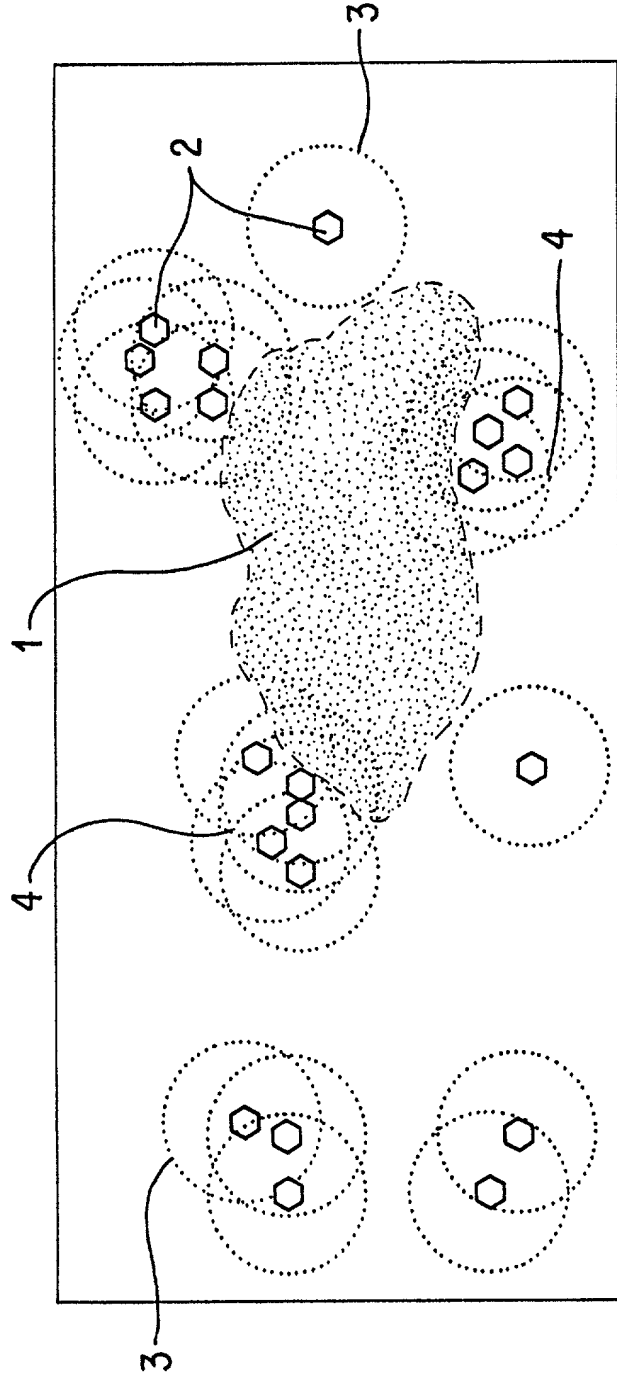


FIG. 1(a)

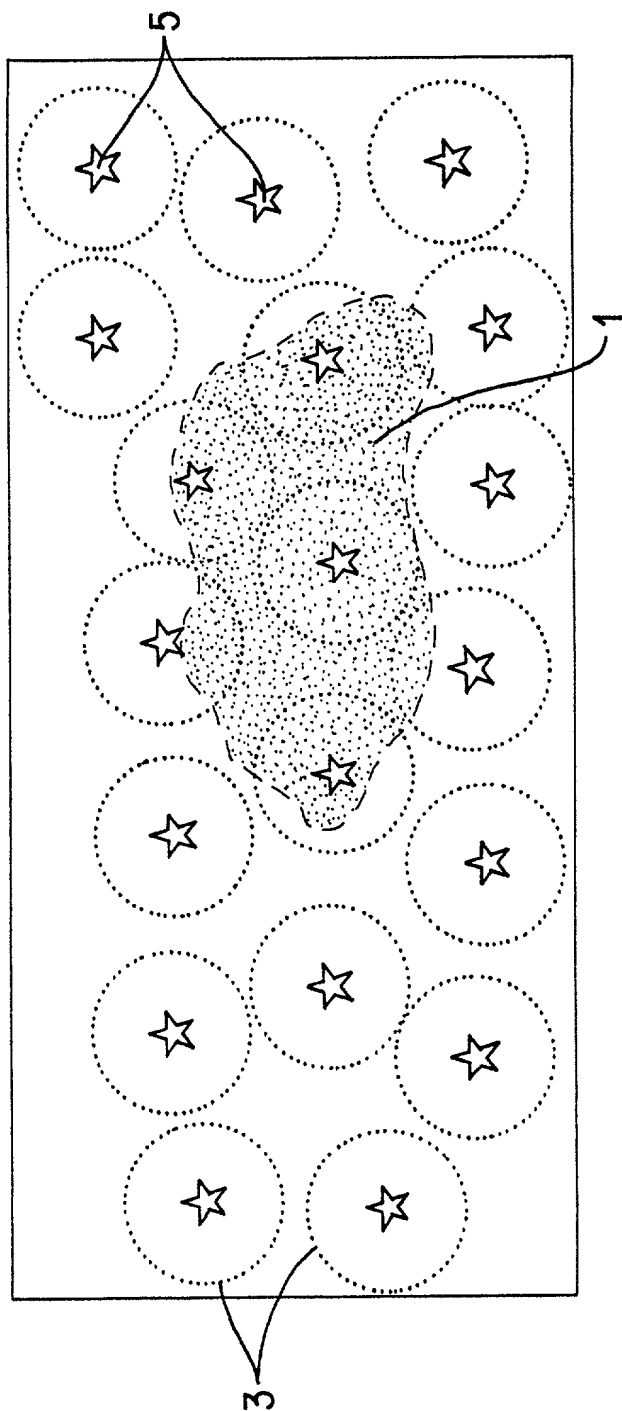


FIG. 1(b)

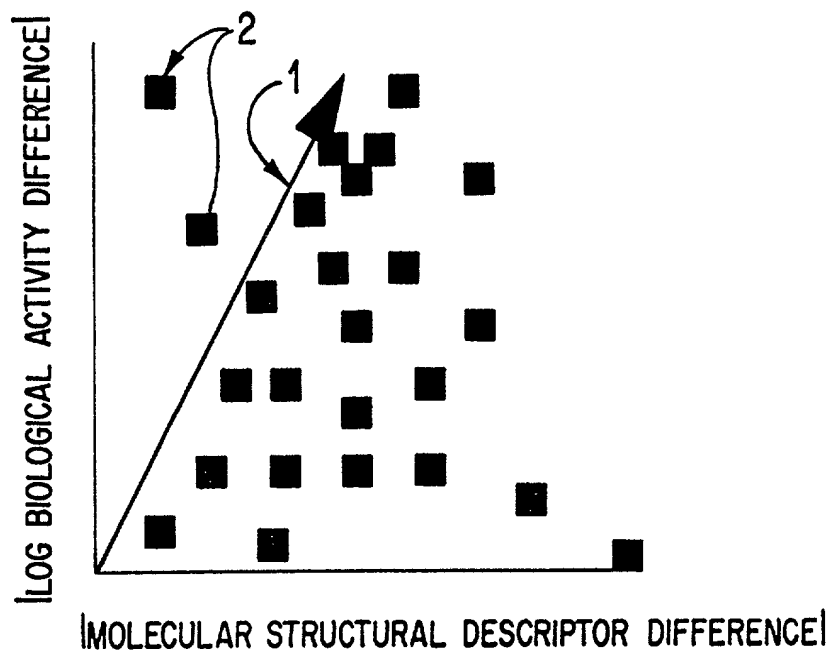


FIG. 2

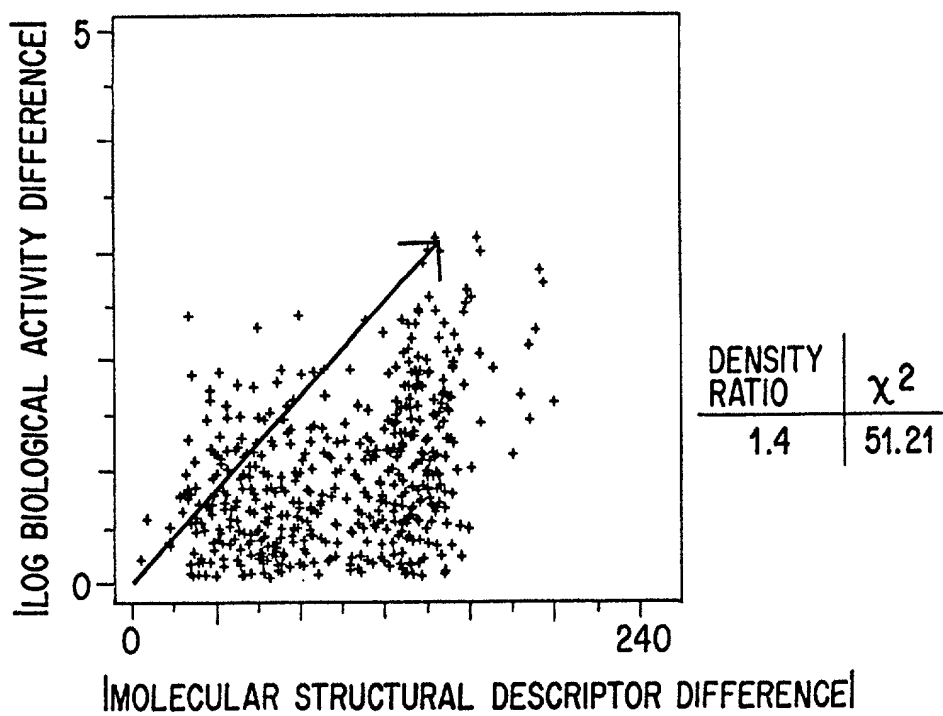


FIG. 3

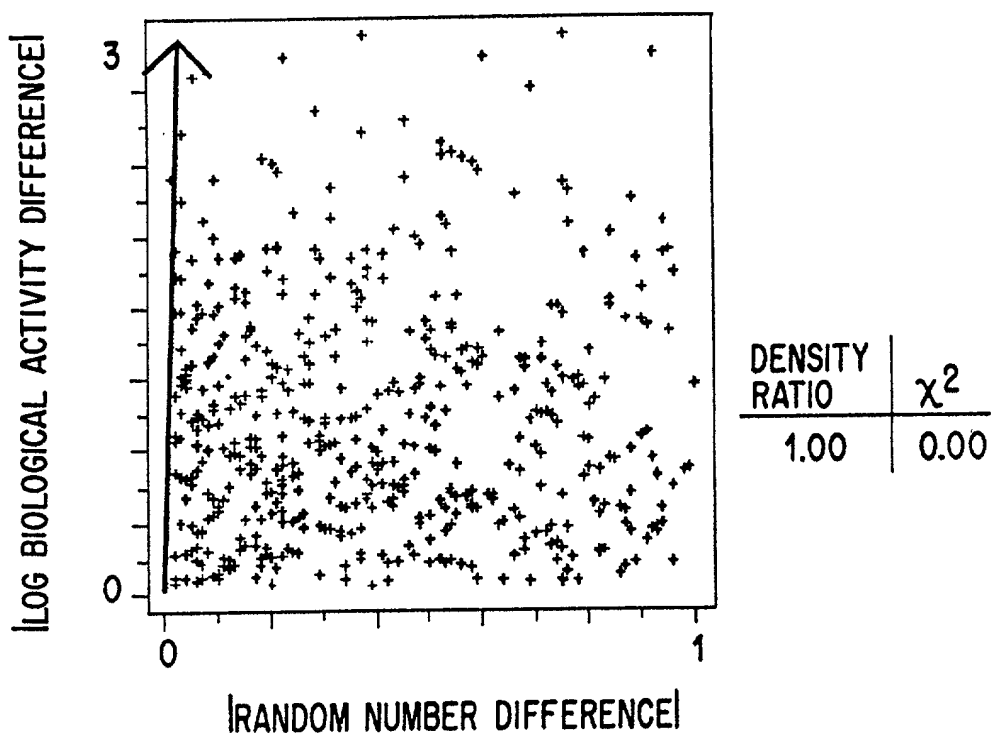


FIG. 4

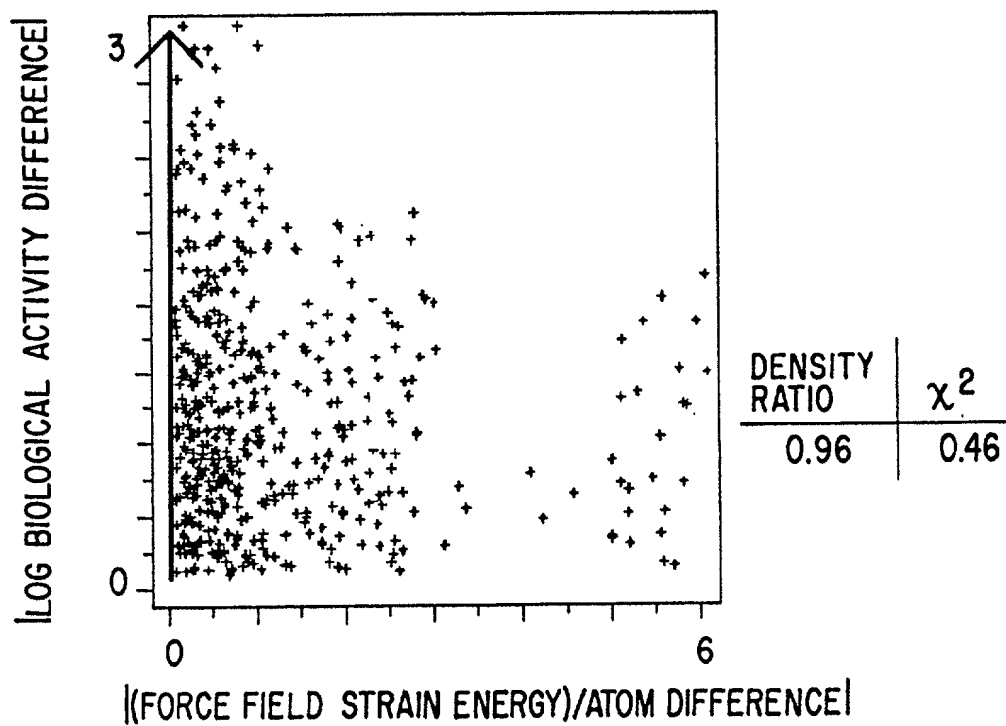
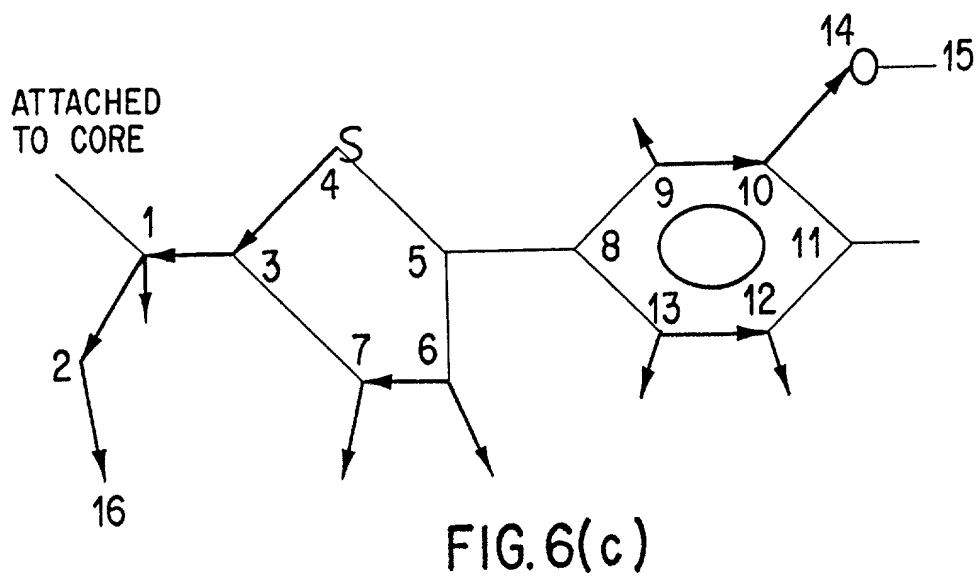
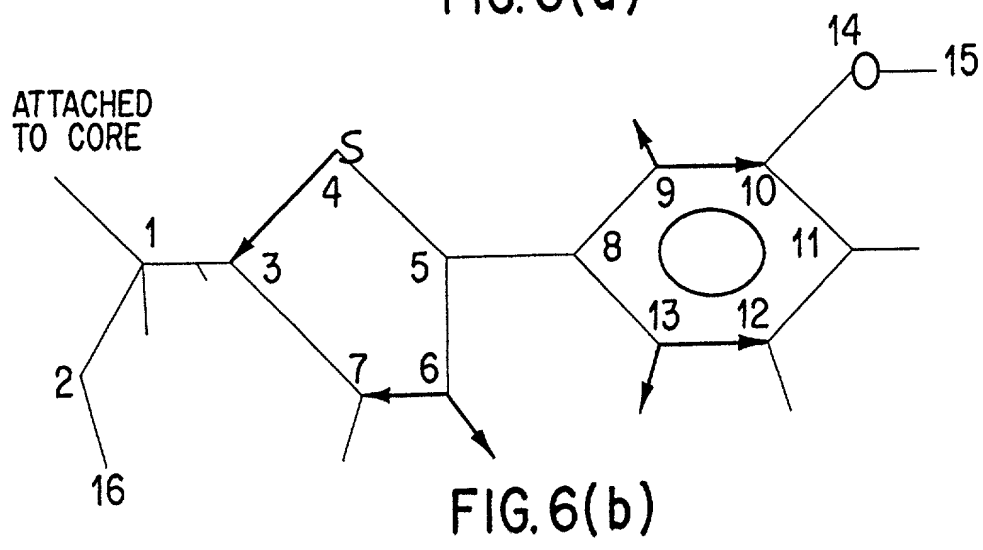
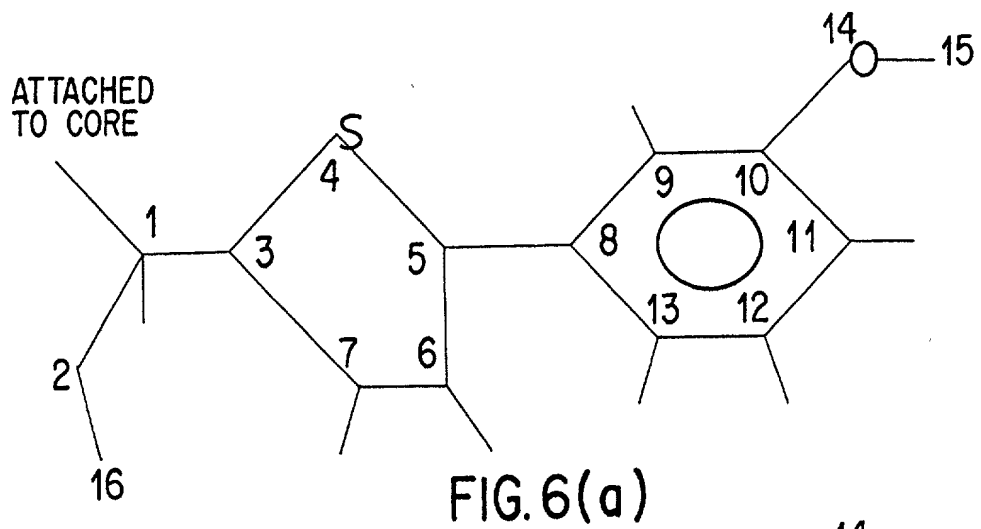


FIG. 5



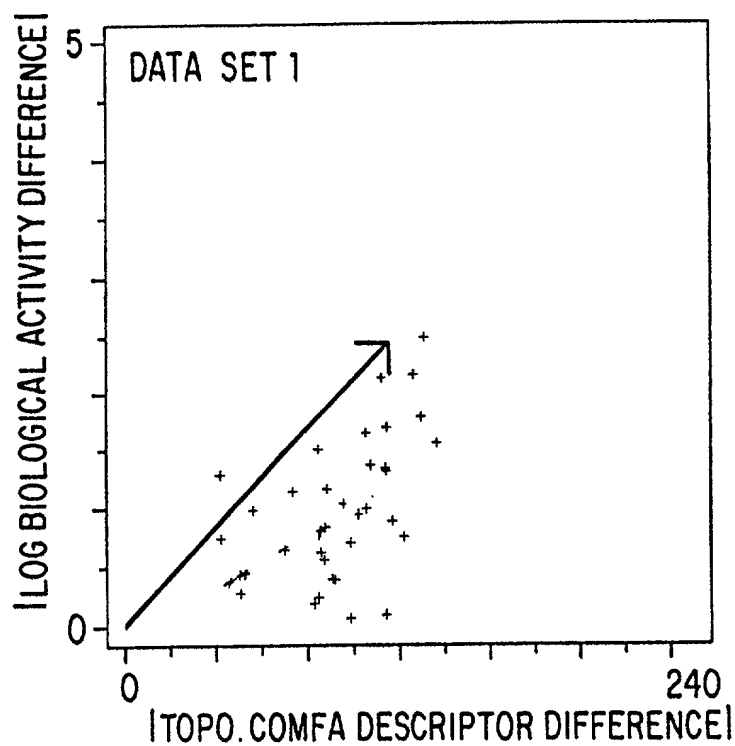


FIG.7(a)

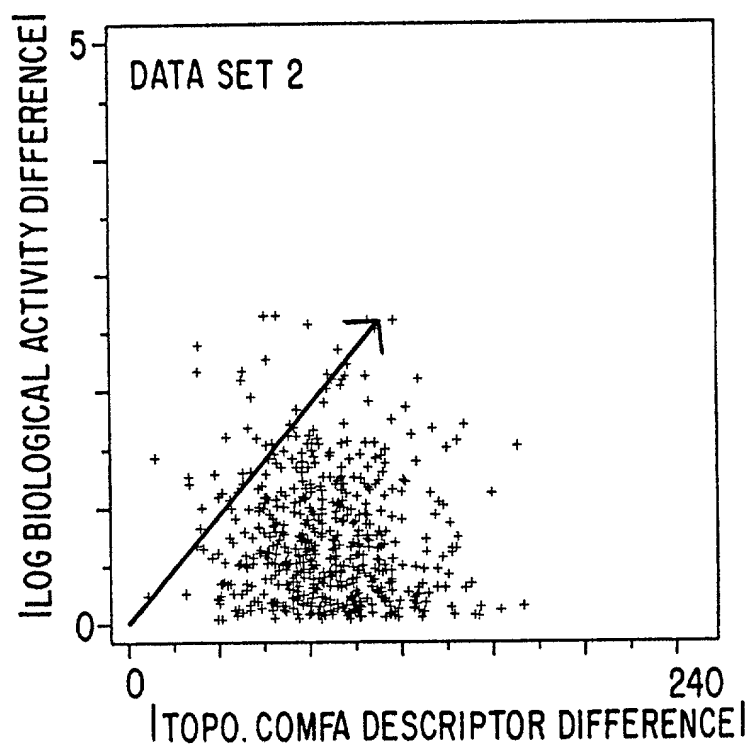


FIG.7(b)

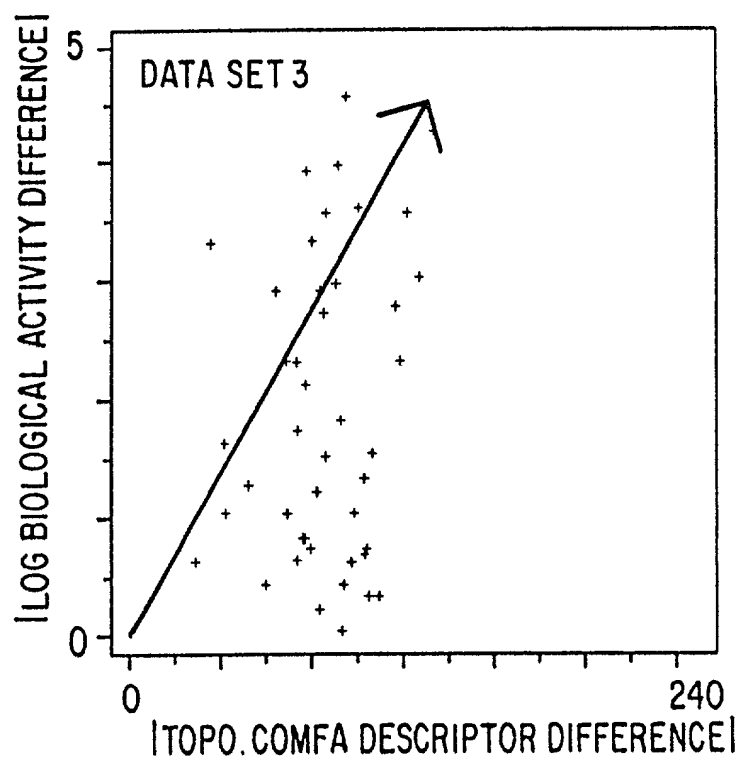


FIG. 7(c)

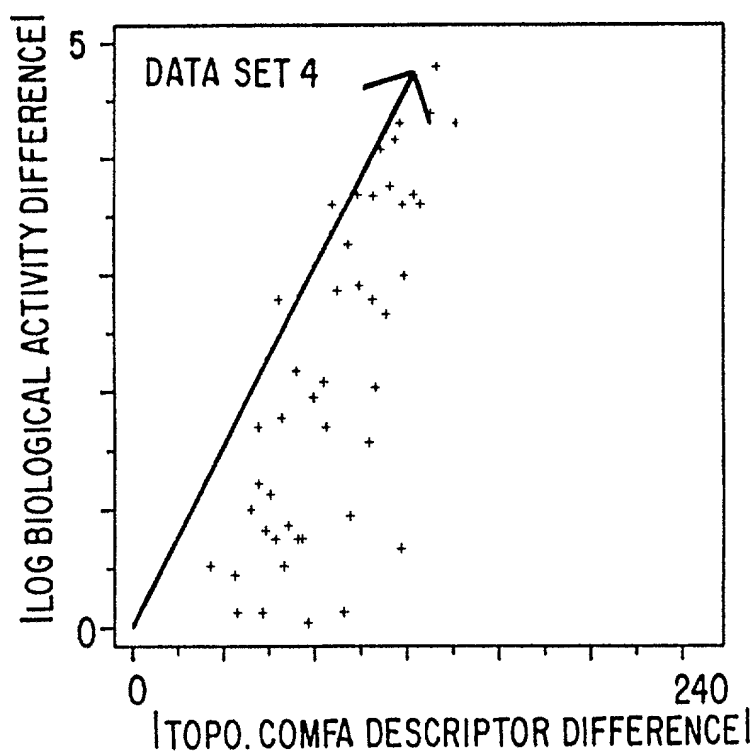


FIG. 7(d)

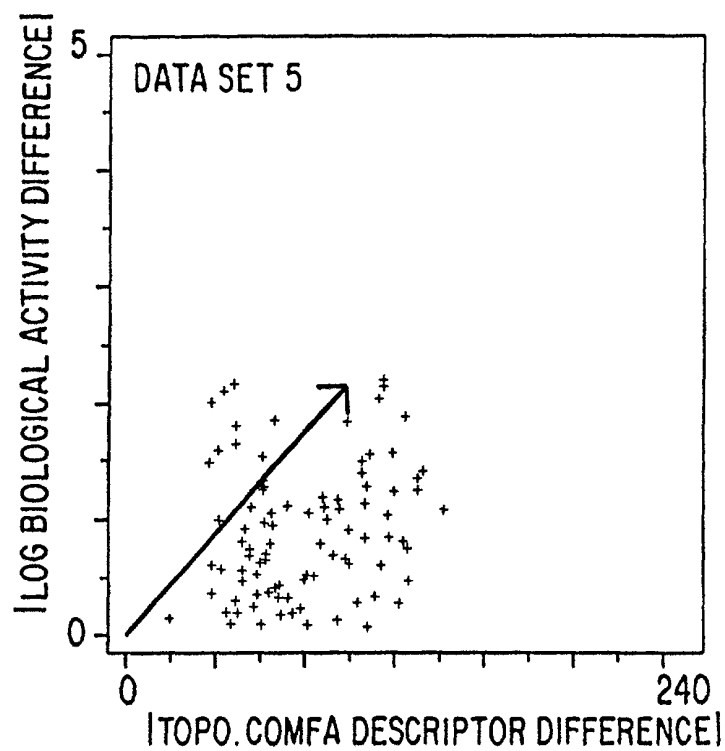


FIG.7(e)

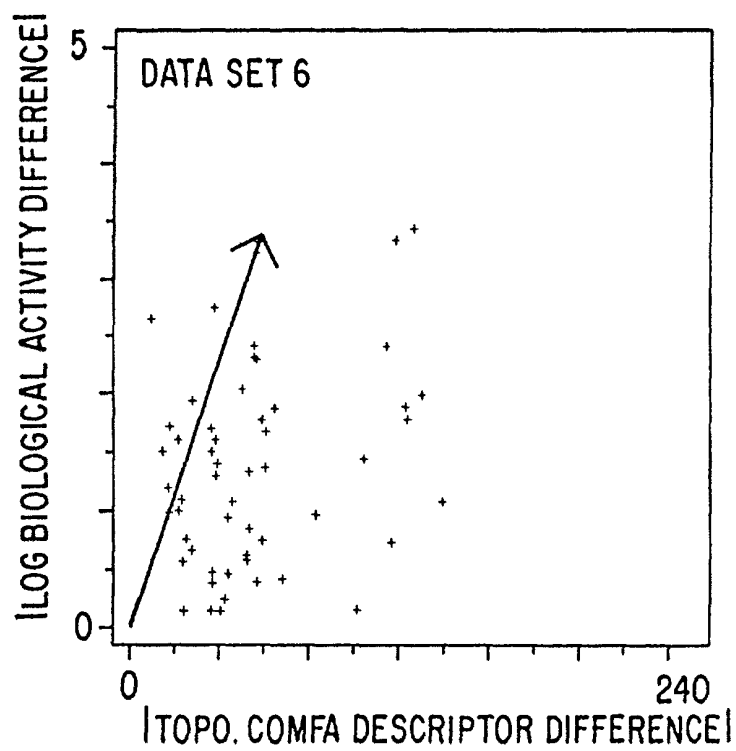


FIG.7(f)

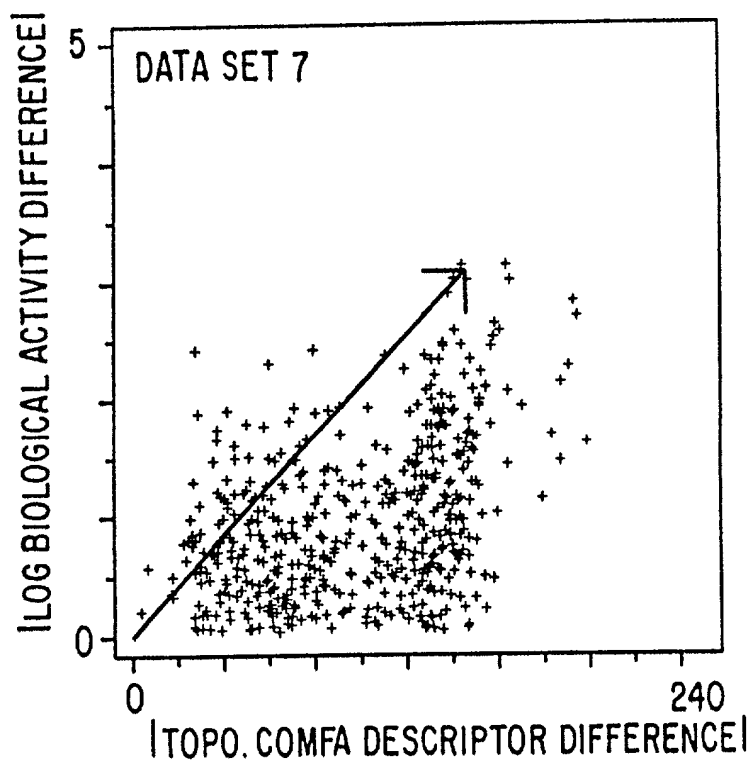


FIG.7(g)

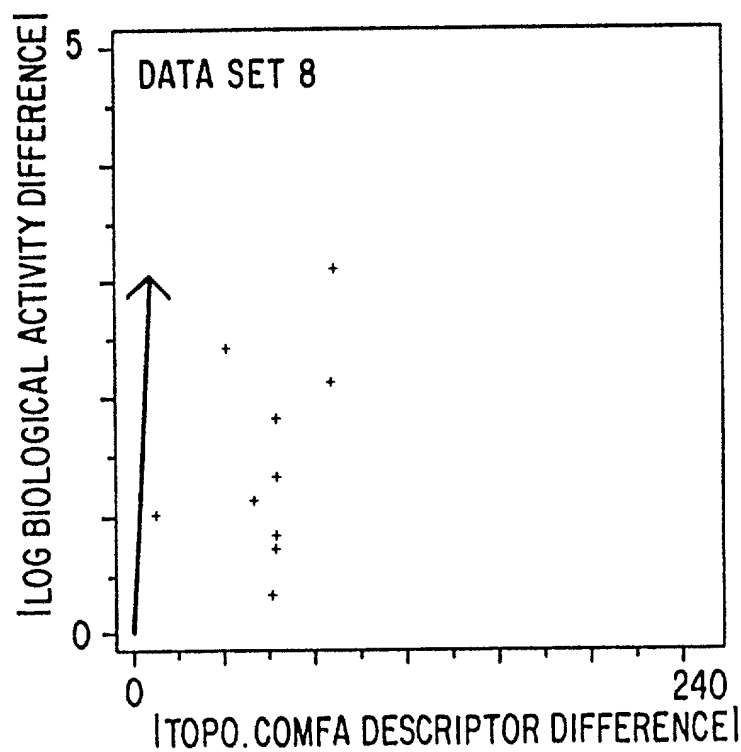


FIG.7(h)

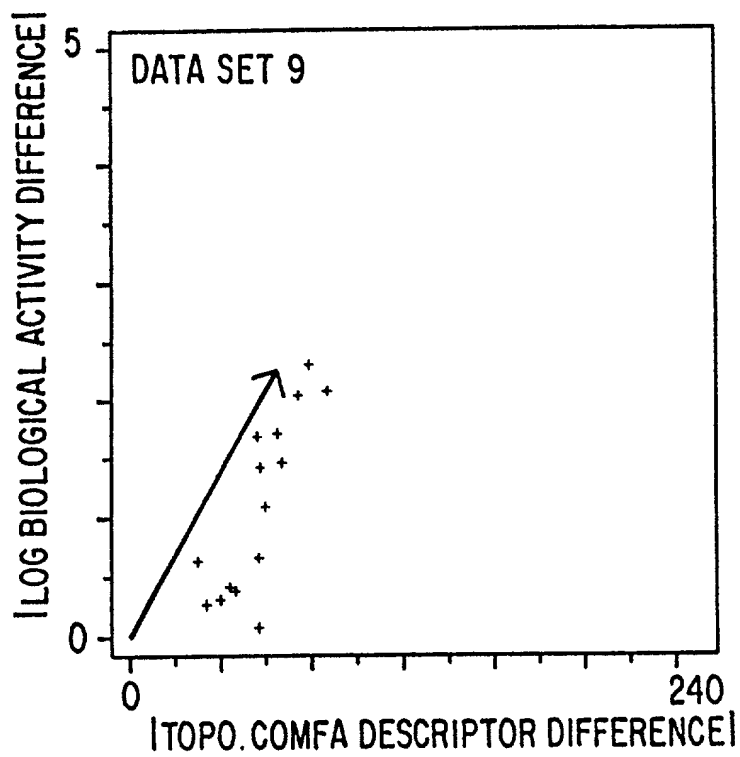


FIG.7(i)

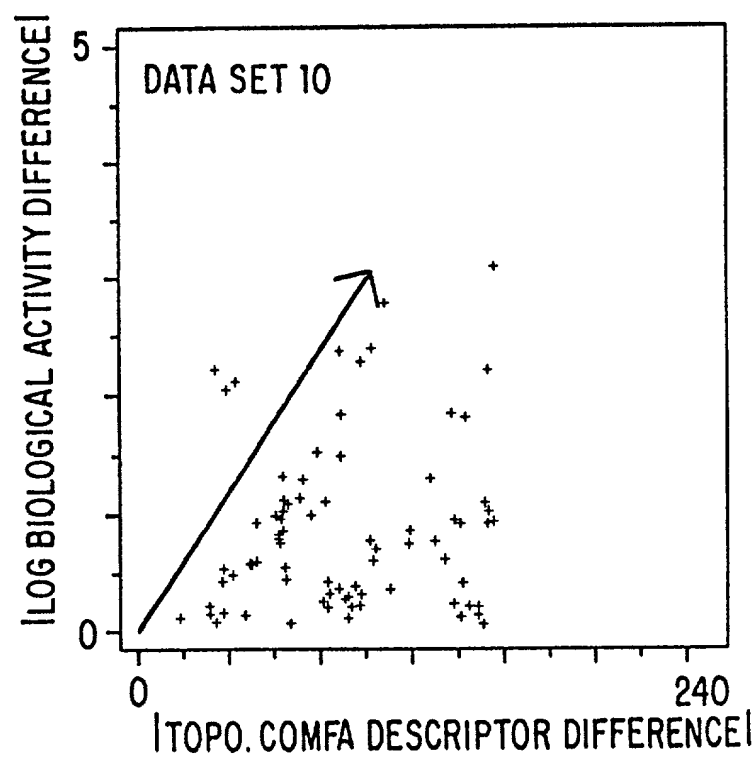


FIG.7(j)

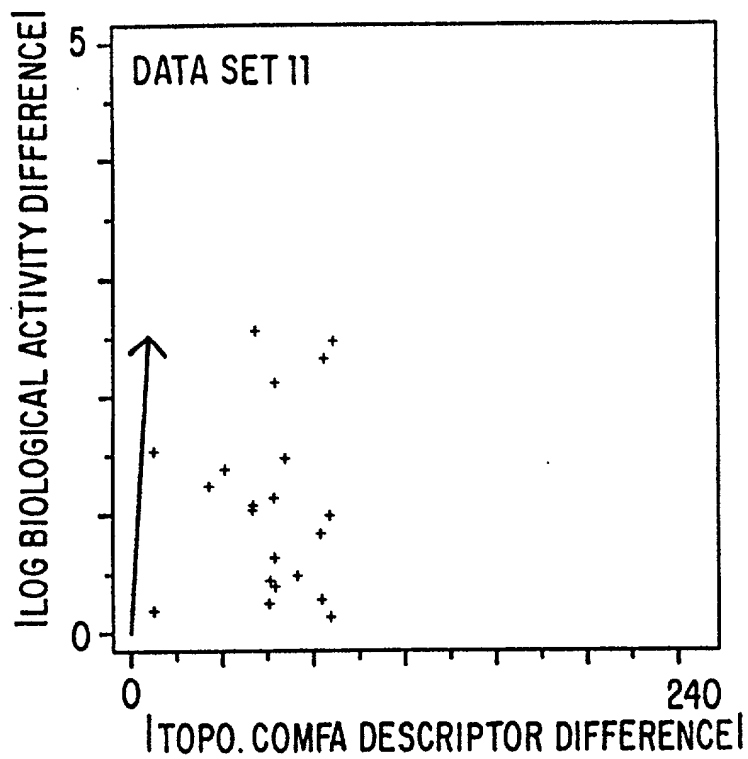


FIG.7(k)

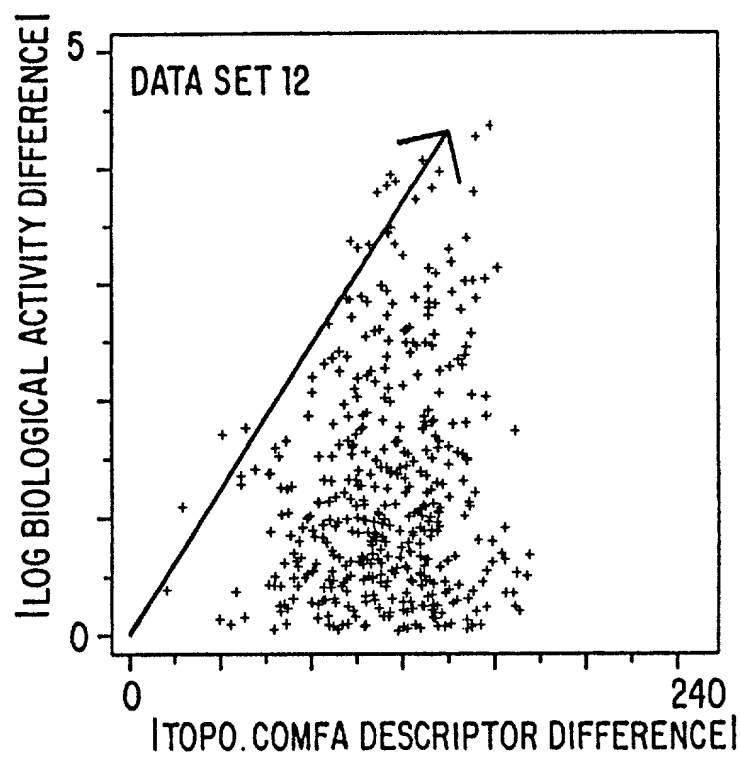


FIG.7(l)

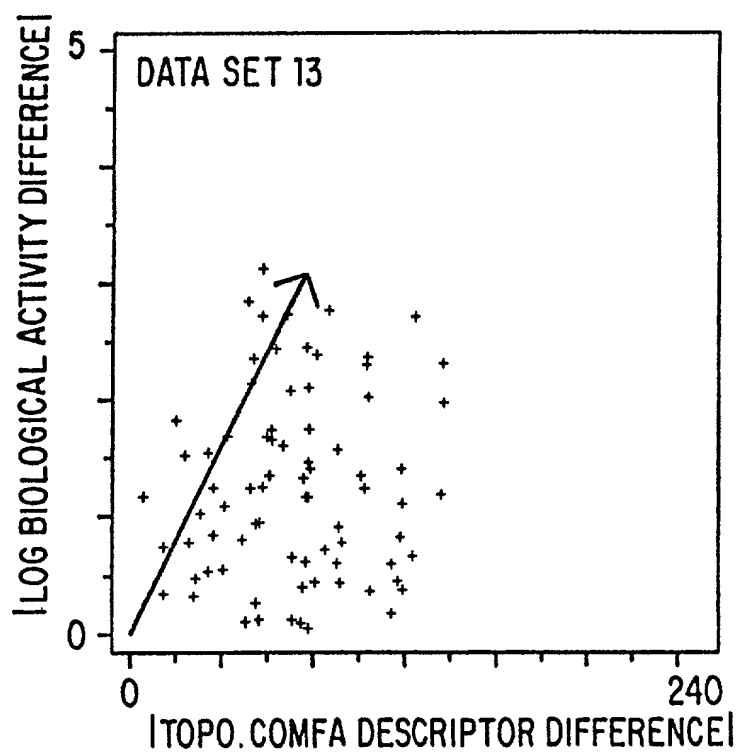


FIG.7(m)

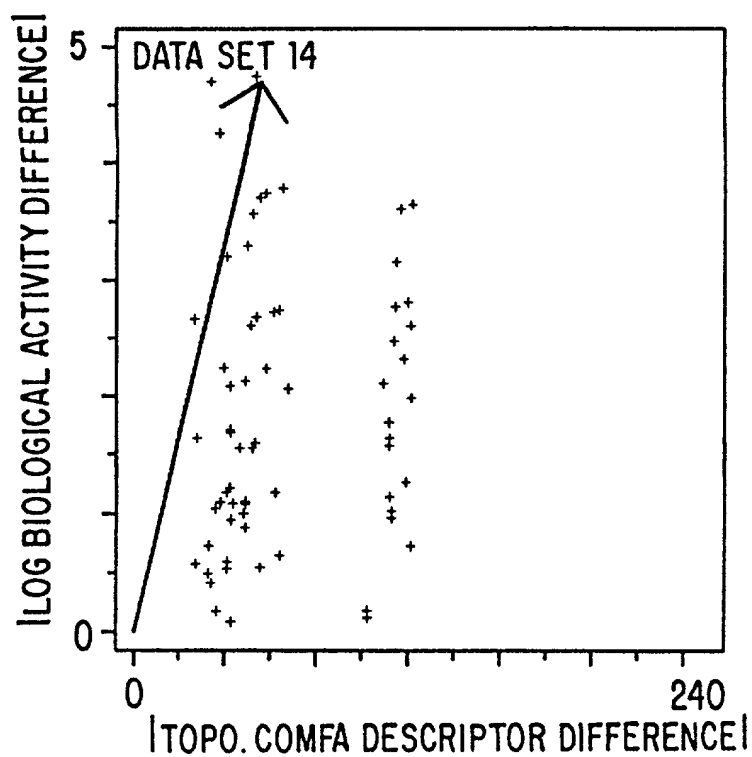


FIG.7(n)

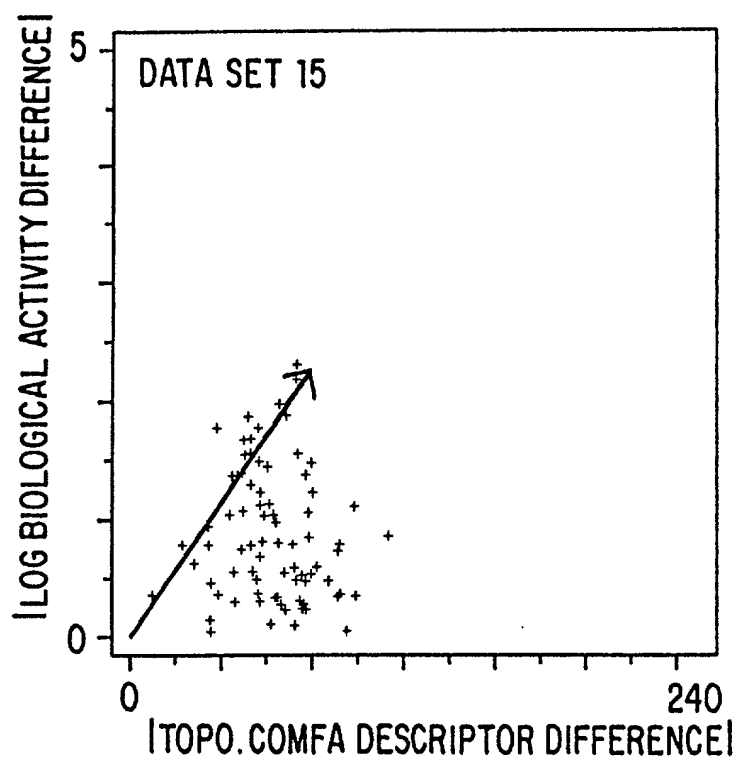


FIG.7(o)

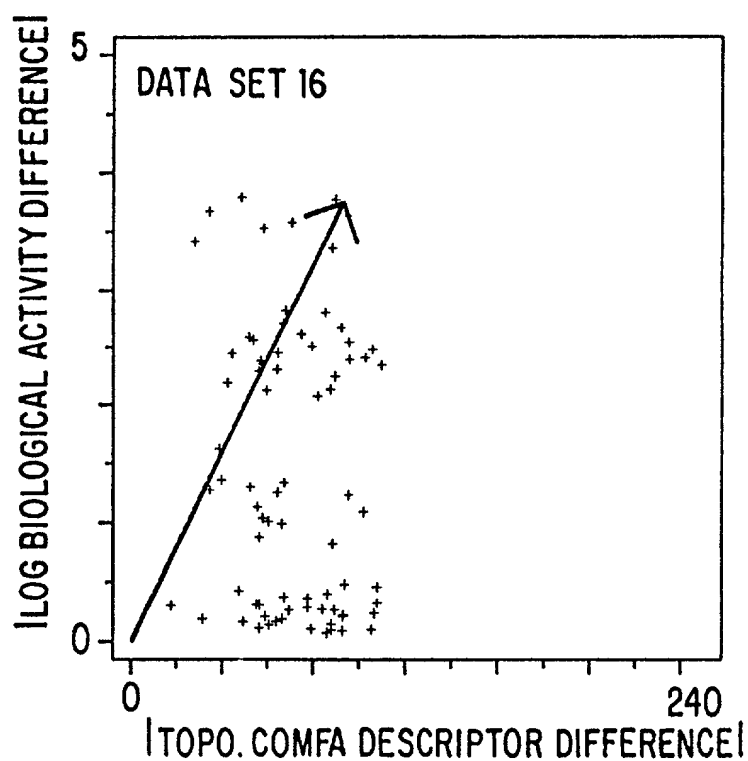


FIG.7(p)

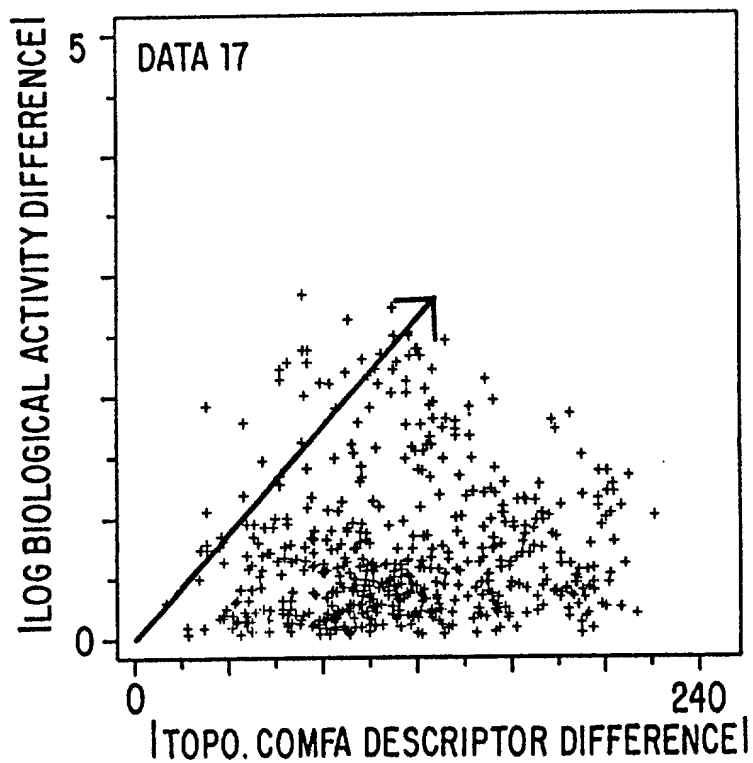


FIG.7(q)

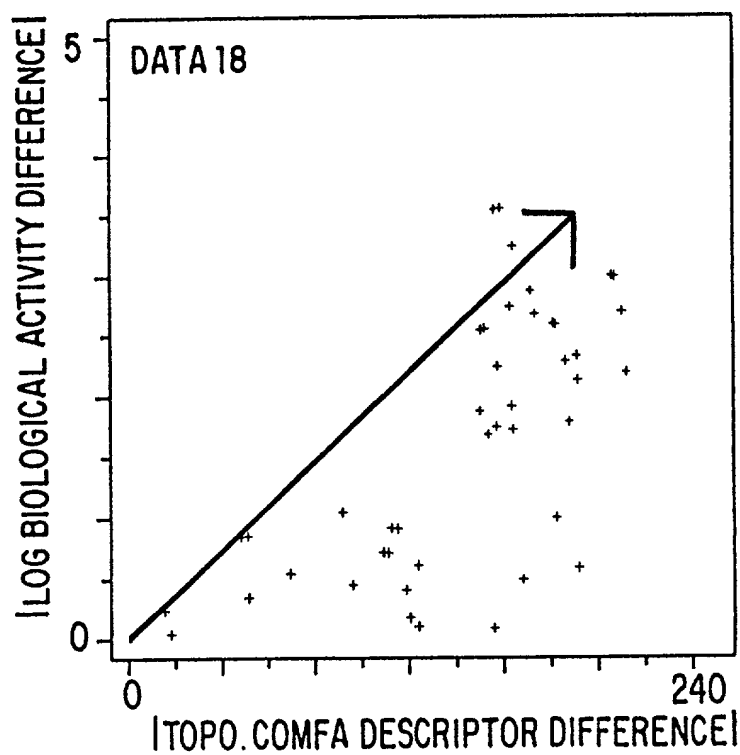


FIG.7(r)

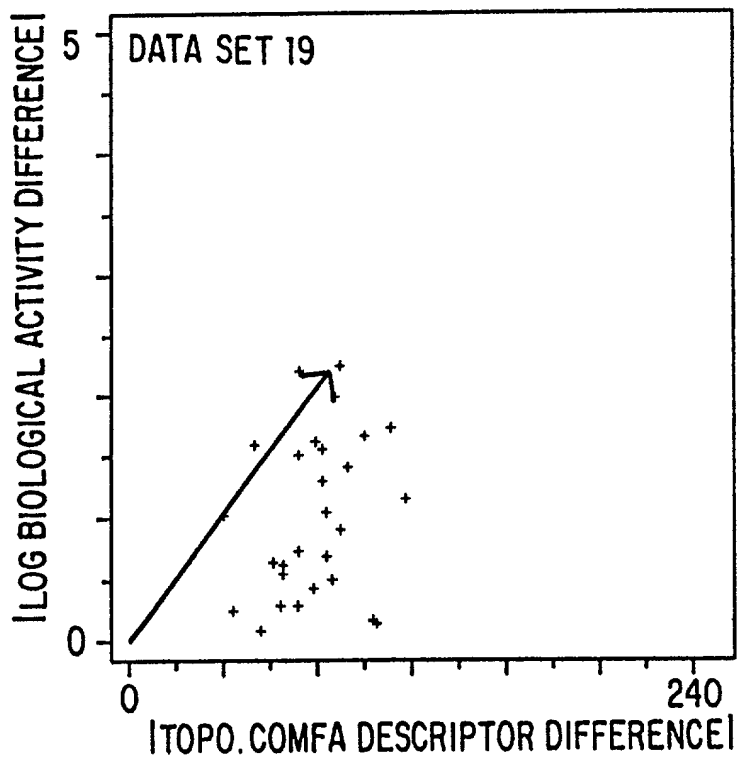


FIG. 7(s)

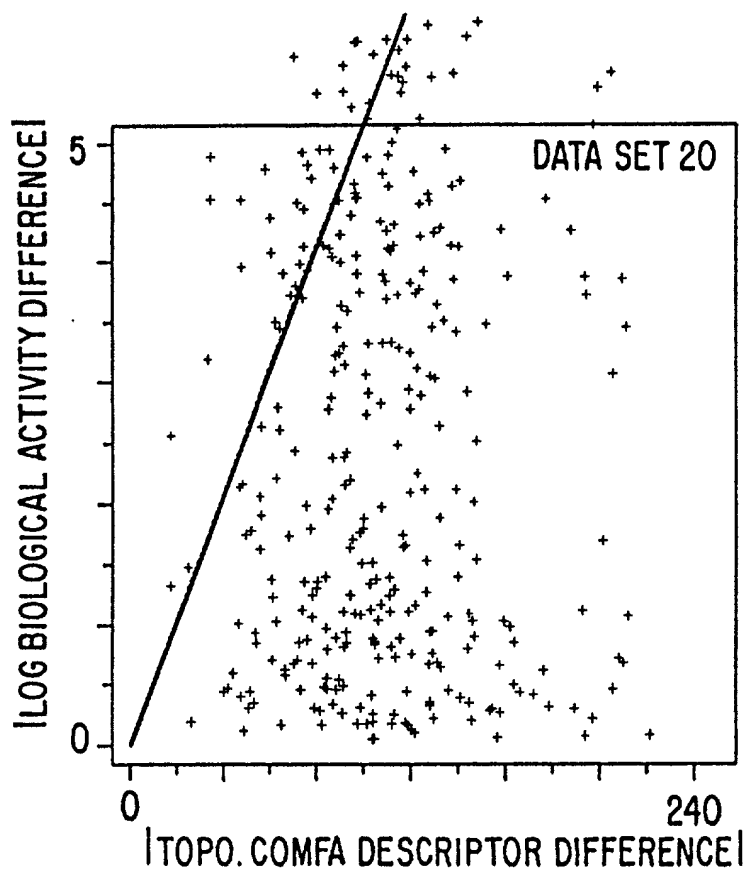


FIG. 7(t)

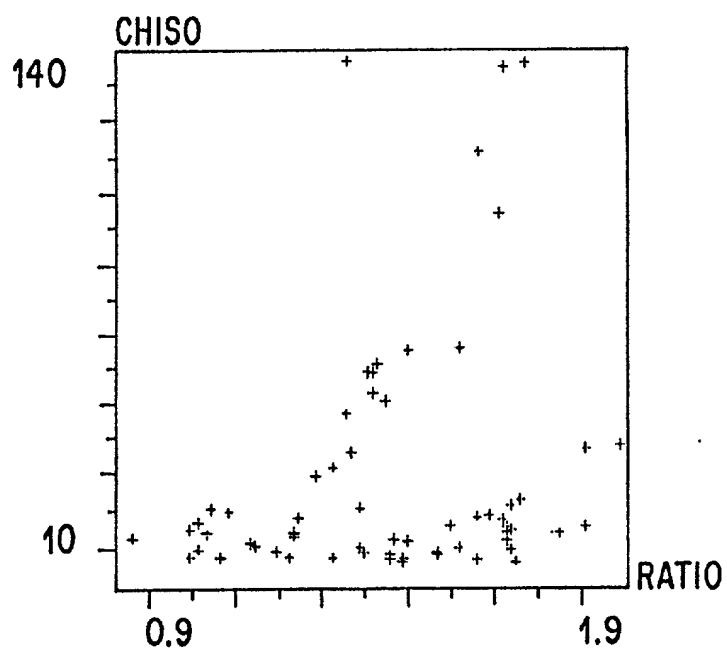


FIG. 8(a)

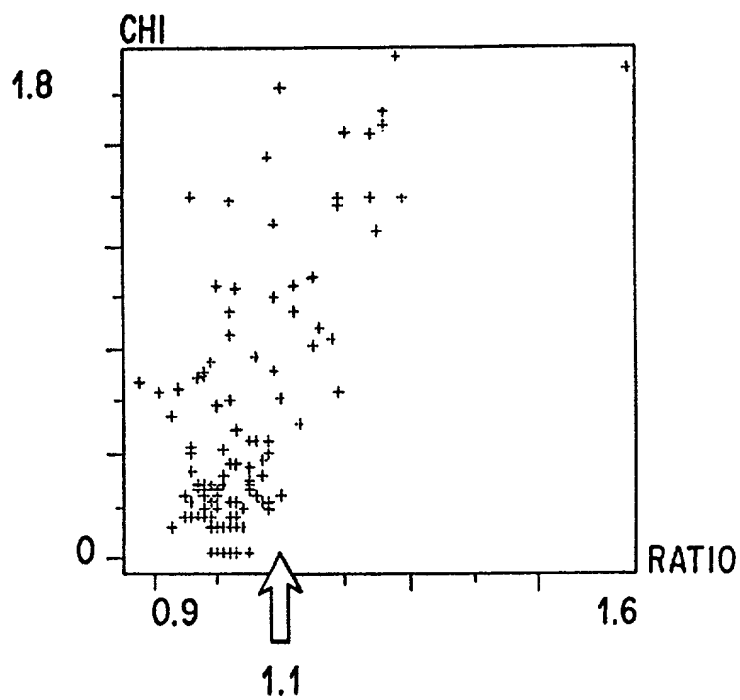


FIG. 8(b)

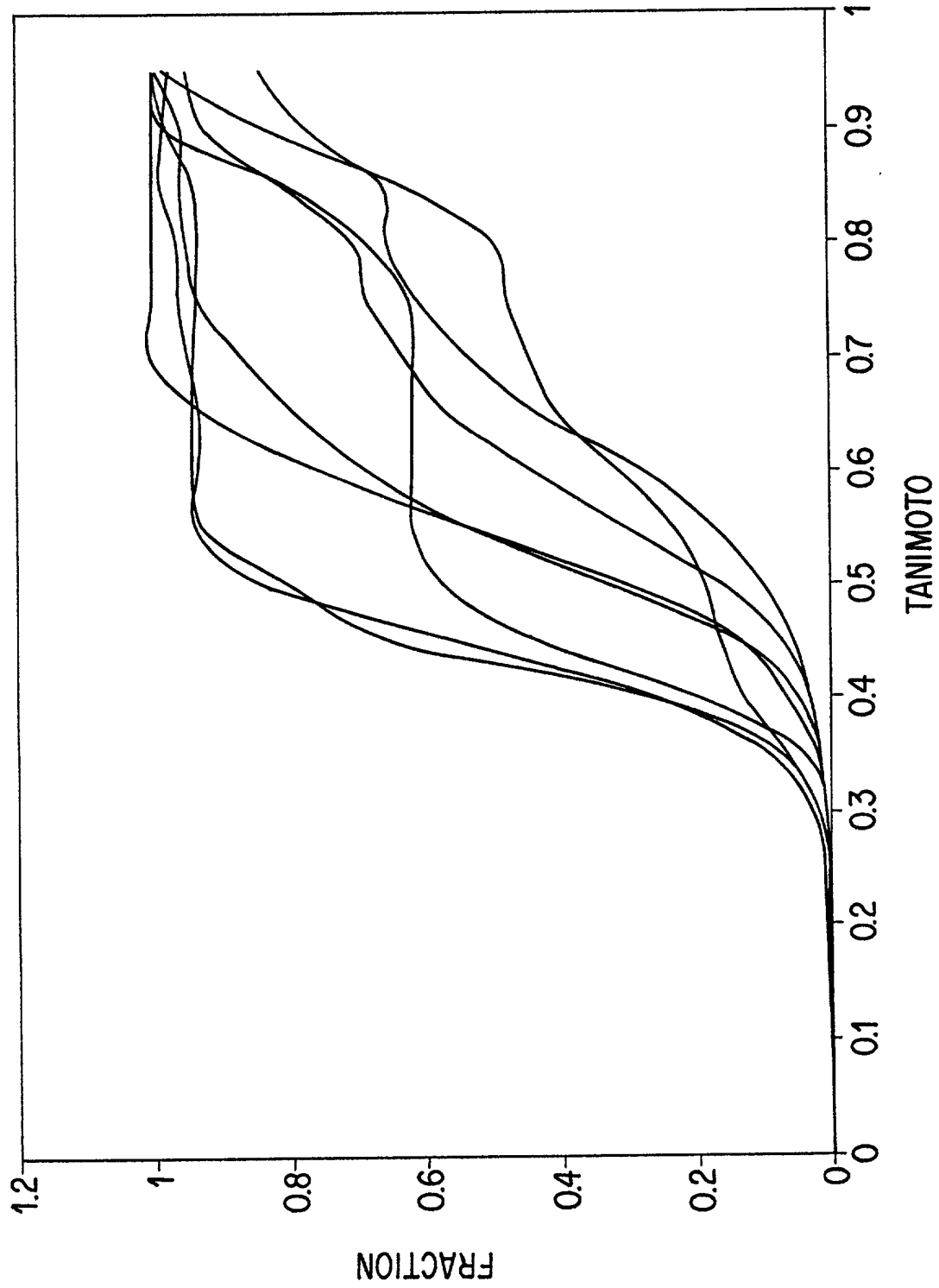


FIG. 9(a)

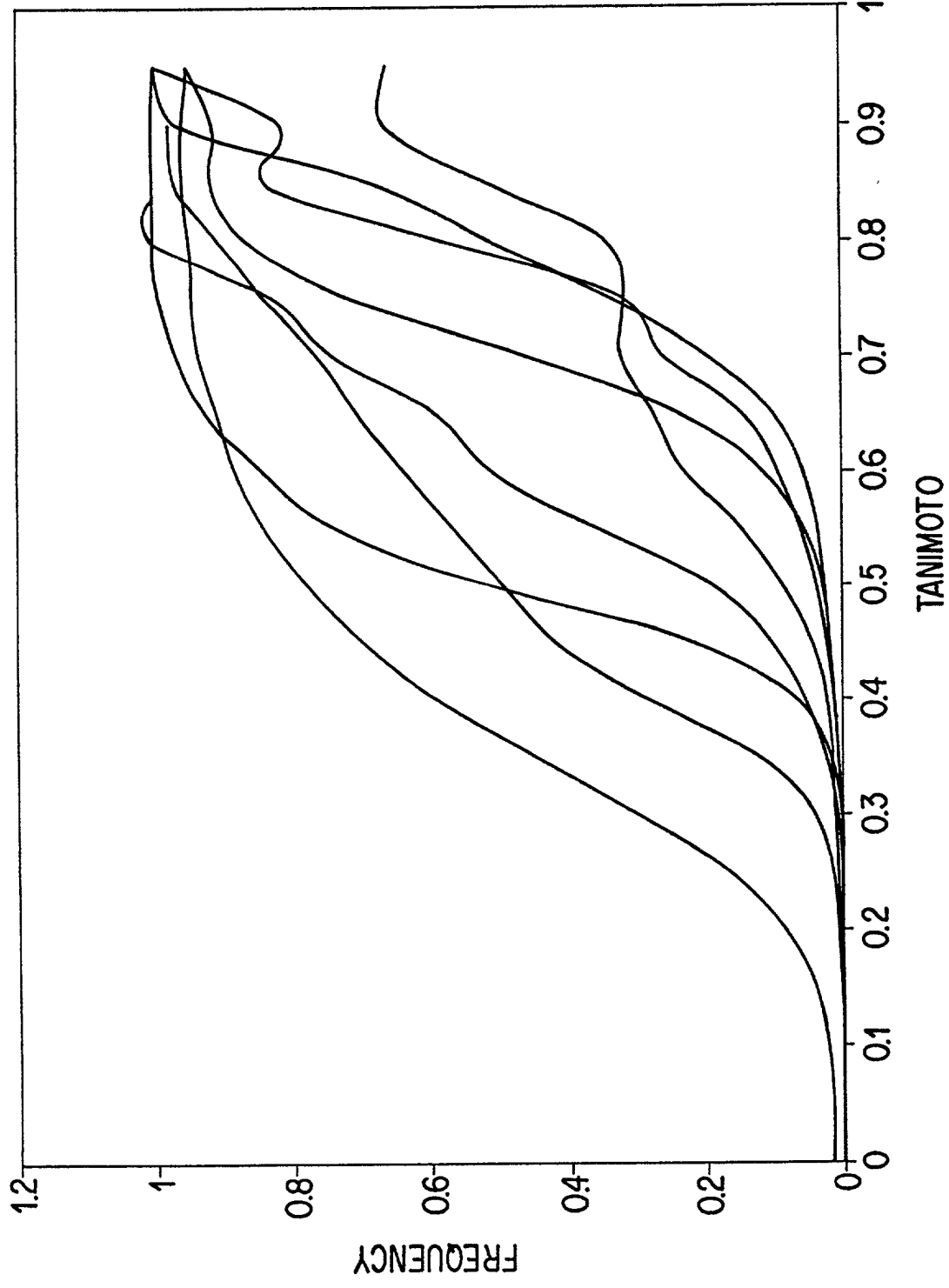
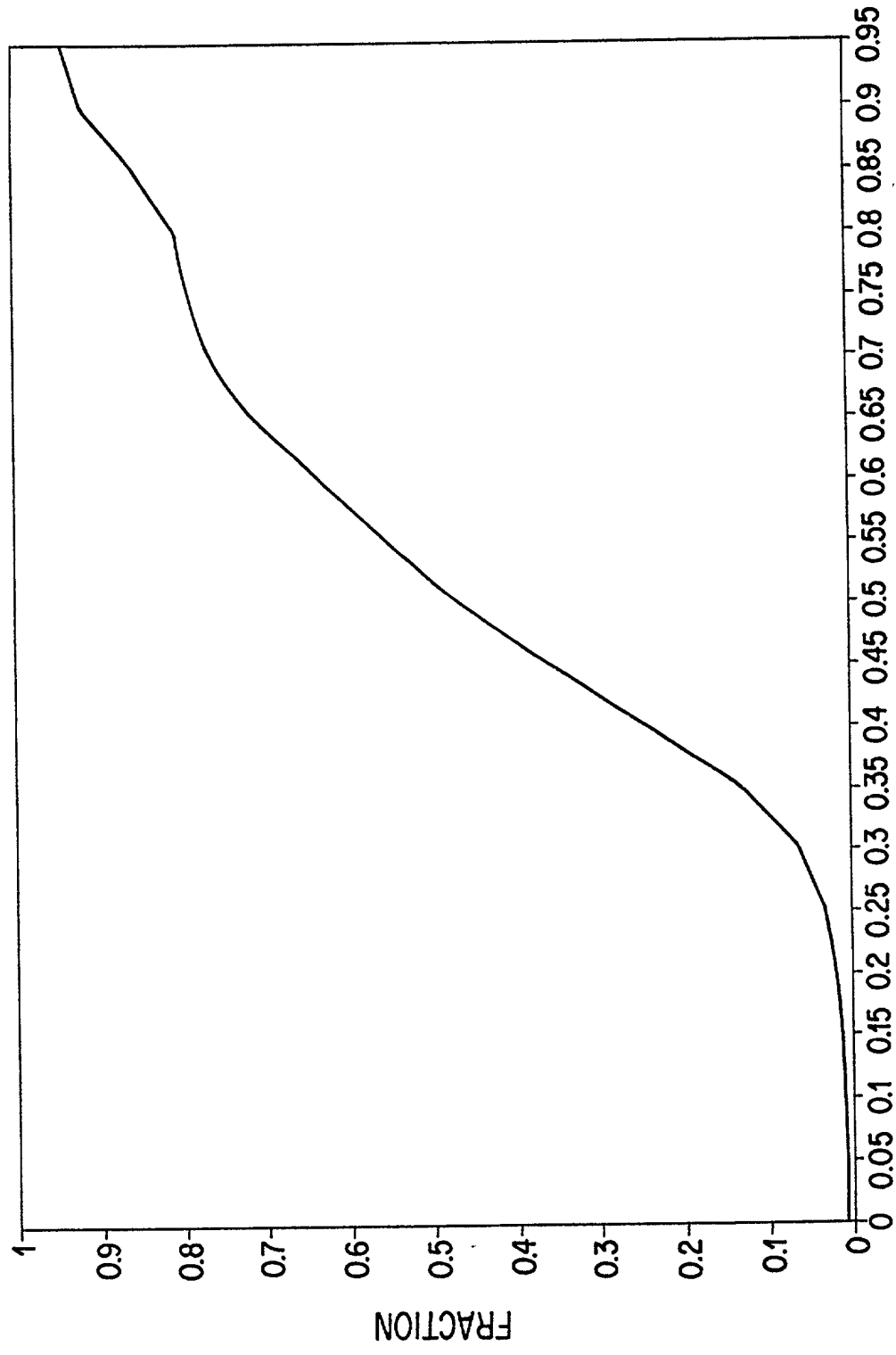


FIG. 9(b)



TANIMOTO

FIG. 9(c)

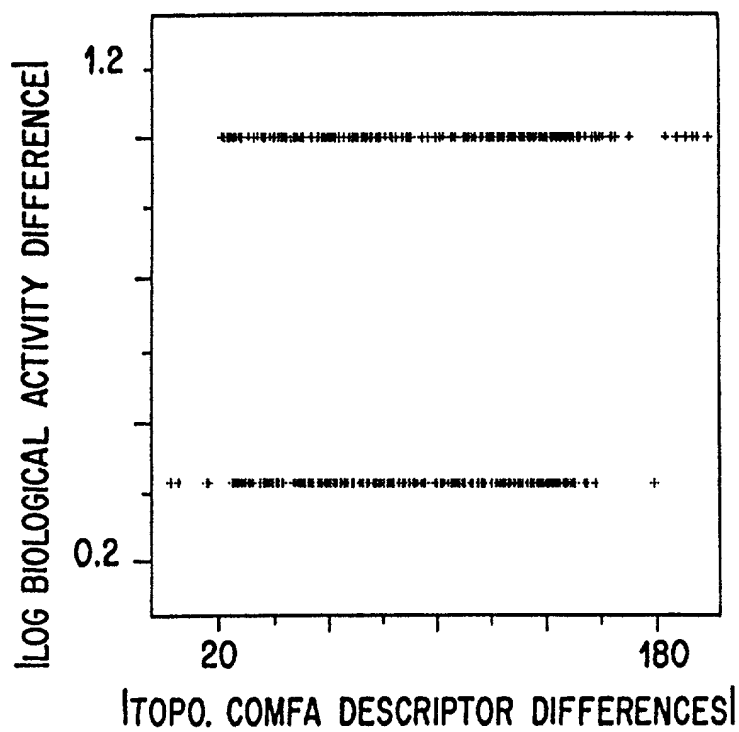


FIG.10(a)

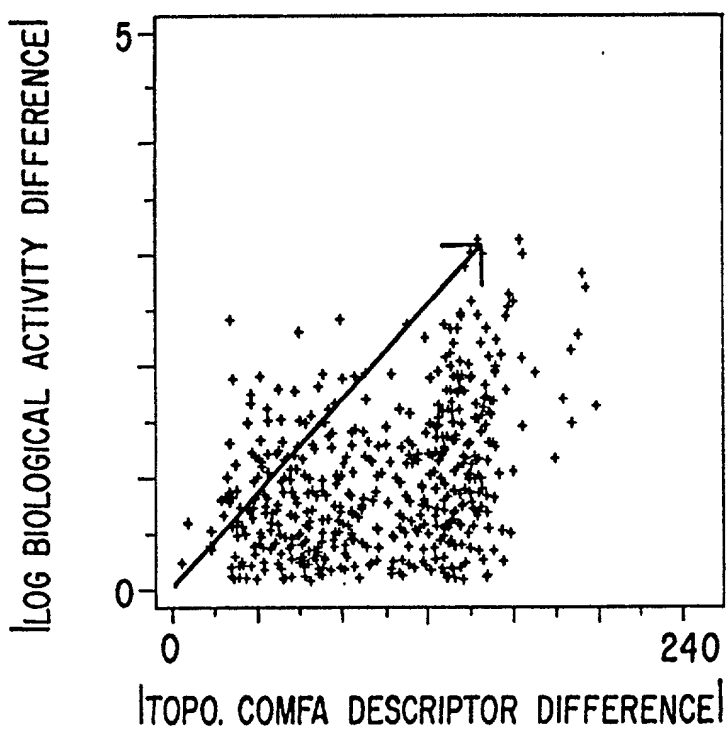


FIG.10(b)

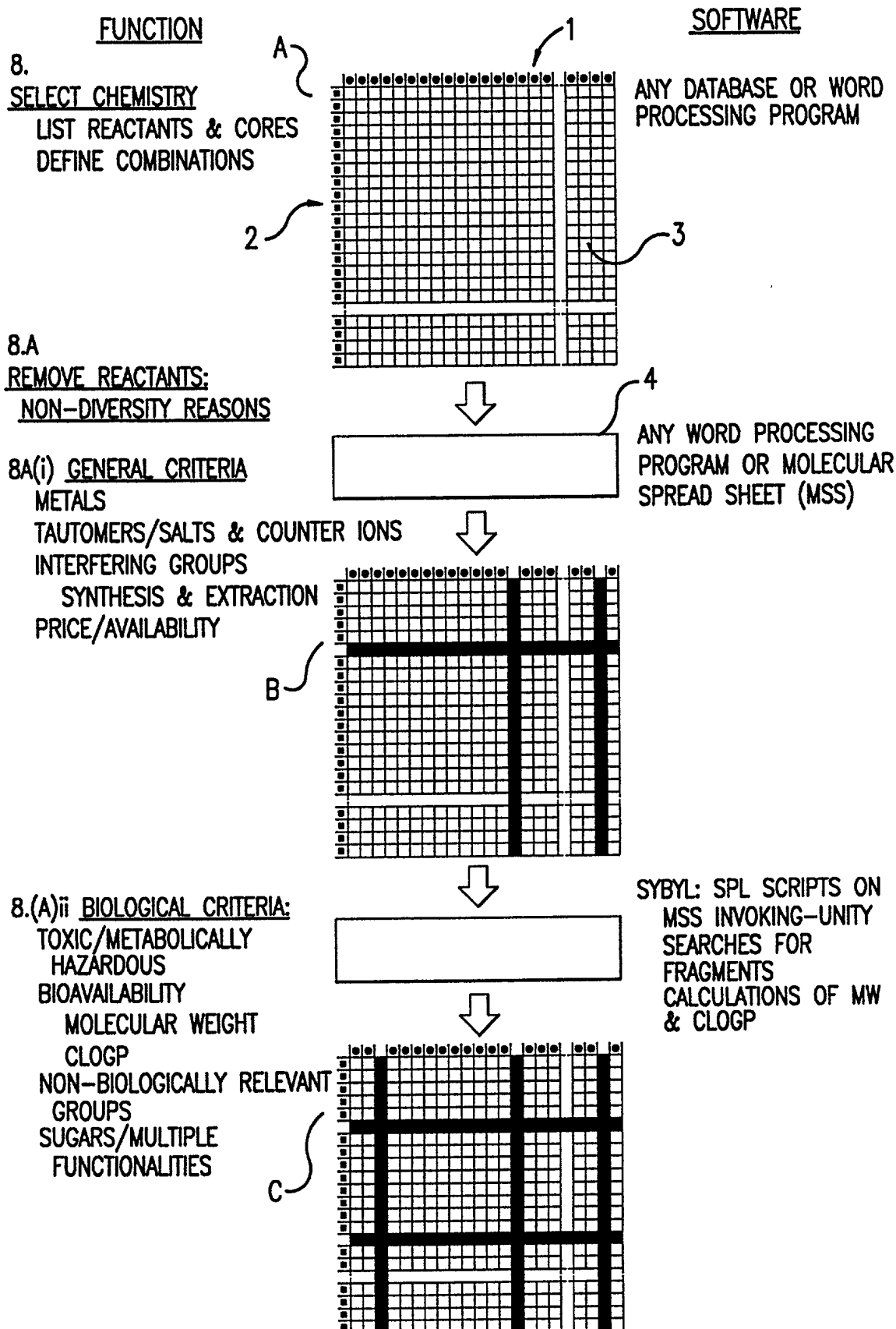


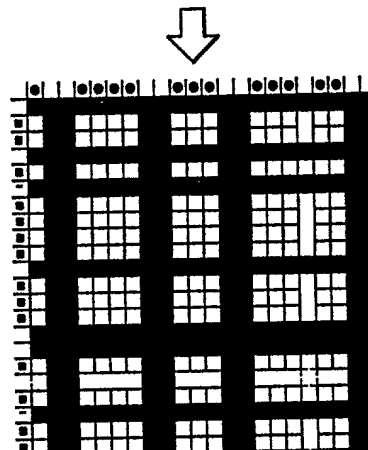
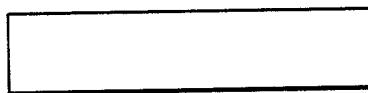
FIG.11(a)

8B

REMOVE NON-DIVERSE REACTANTS:

3D STRUCTURE GENERATION
 TOPOMERIC CONFORMER
 ALIGNMENT
 CoMFA FIELD GENERATION:
 HYDROGEN BOND FIELD
 GENERATION
 ROTATABLE BOND FIELD
 ATTENUATION
 CALCULATE FIELD
 DIFFERENCES
 CLUSTER
 REACTANT SELECTION FROM
 EACH CLUSTER

D



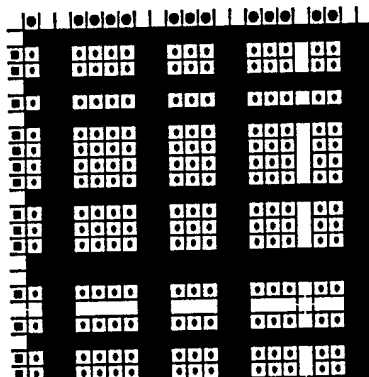
SYBYL: SELECTOR &
 CONCORD 3.2.1
 SPECIALIZED SOFTWARE:
 APPENDIX "A"
 (SYBYL: CoMFA
 STERIC & MSS)

SYBYL: HIERCHICAL
 CLUSTER - MSS
 SYLBL: MSS

8C

COMBINE REACTANTS TO BUILD PRODUCTS

E



SYBYL: LEGION MSS

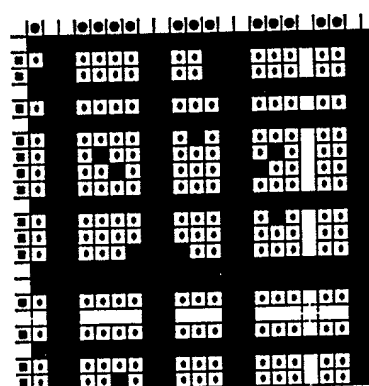
8D

REMOVE PRODUCTS:

NON-DIVERSITY REASONS

GENERAL CRITERIA:

F



SYBYL: SPL SCRIPTS
 ON MSS

FIG.11(b)

8D

BIOLOGICAL CRITERIA:

METABOLICALLY HAZARDOUS

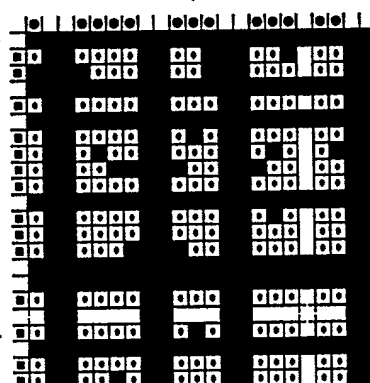
BIOAVAILABILITY

MOLECULAR WEIGHT

CLOGP



G



SYBYL: SPL SCRIPTS ON
MSS INVOKING -

UNITY SEARCHES FOR
FRAGMENTS

CALCULATIONS OF MW
& CLOGP

8E

REMOVE NON-DIVERSE
PRODUCTS:

GENERATE TANIMOTO

2D FINGERPRINTS

SAMPLE PRODUCTS:

EXCLUDE TAN ≥ 0.85



UNITY



H

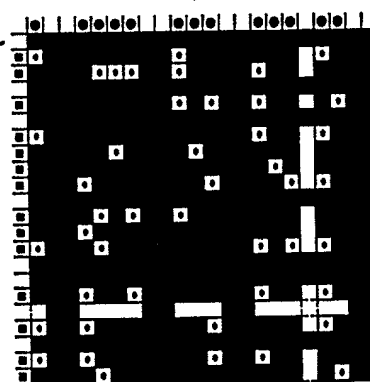
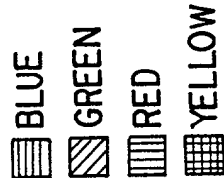


FIG.11(c)



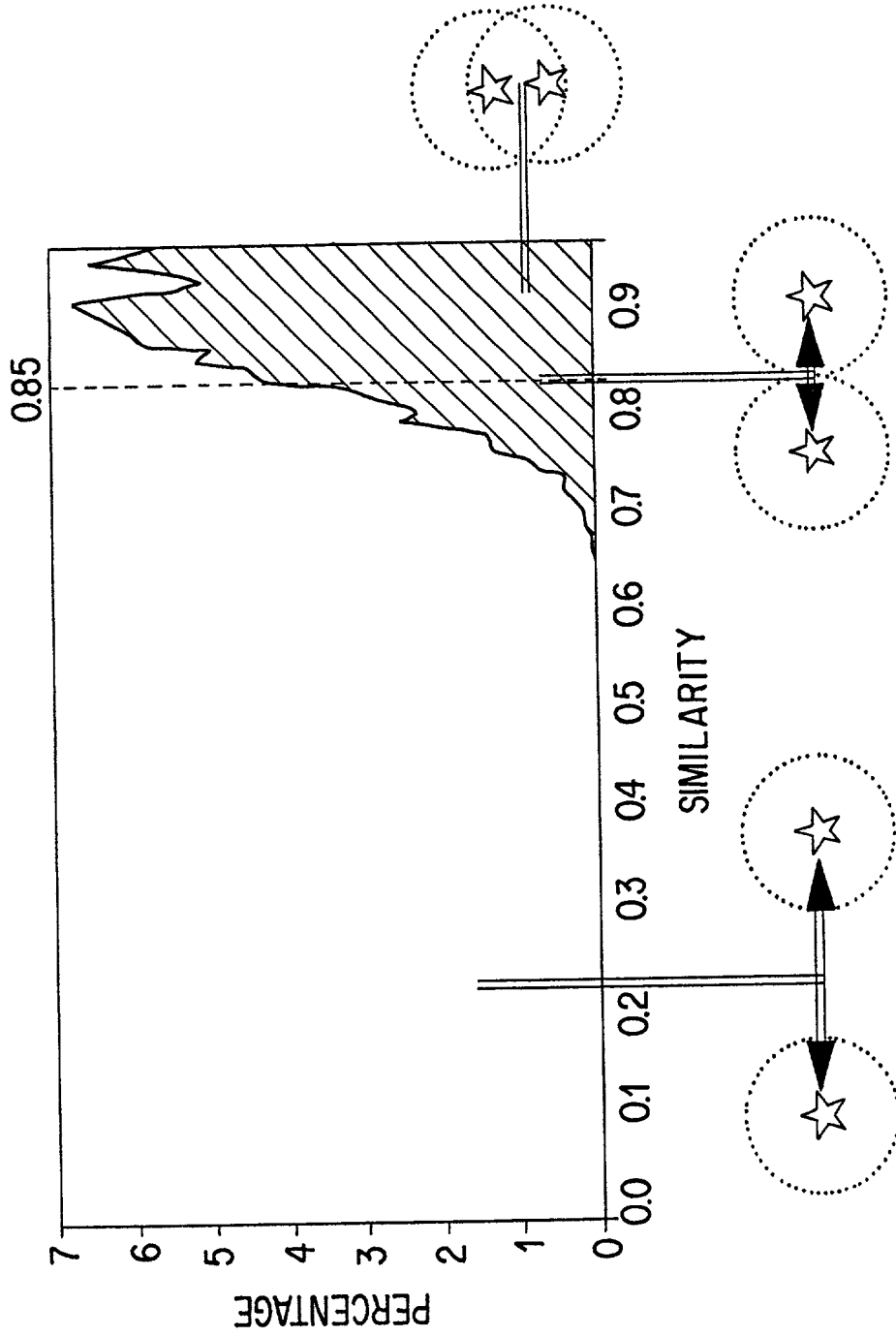


FIG. 13

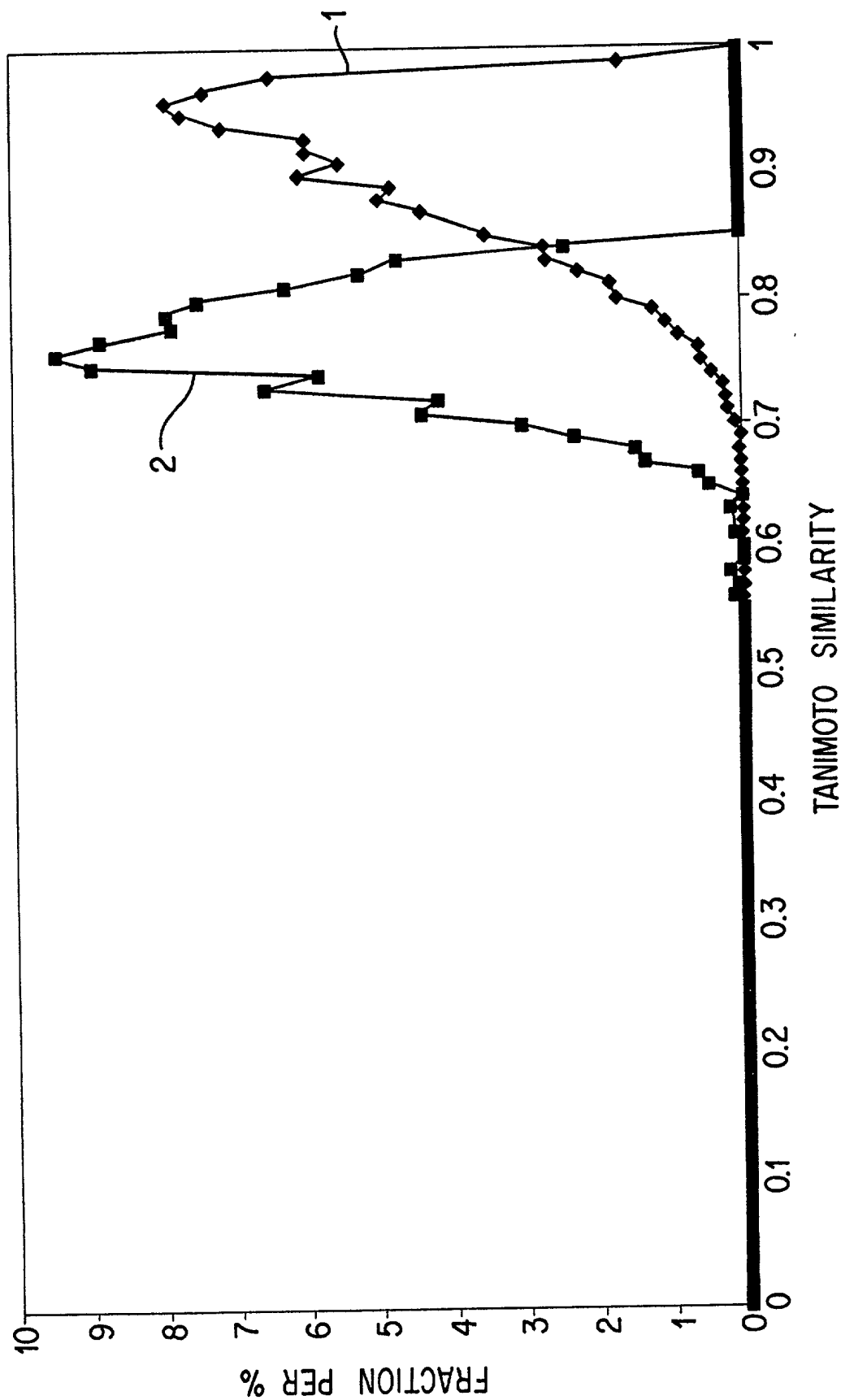


FIG. 14

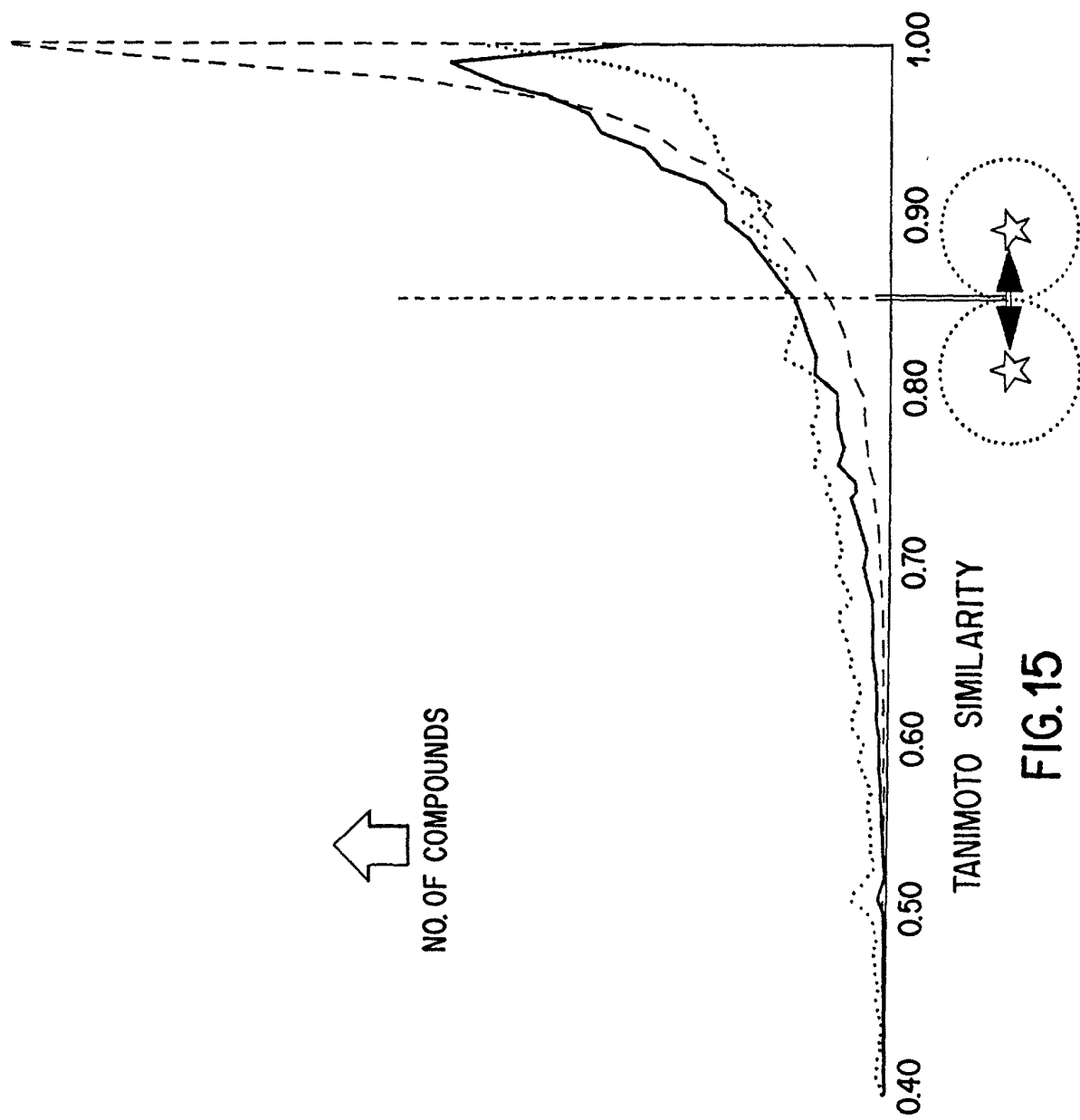


FIG. 15